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DOCUMENT DESCRIPTION	DOCUMENT CONTROL NUMBER	DATE RECEIVED
8EHQ-11- 18413	88110000297	8/19/11

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August 18, 2011

Via Federal Express

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Room 6428
Attention: 8(e) Coordinator
Office of Pollution Prevention and Toxics
U.S. Environmental Protection Agency
1201 Constitution Ave., NW
Washington, DC 20004

Dear 8(e) Coordinator:

6-Ethoxy-N-methyl-l,3,5-triazine-2,4-diamine
CAS# 62096-63-3

This letter is to inform you of the results of a 28-day oral toxicity study and a biodegradability test with the above referenced test substance.

28-day oral study

The test substance, formulated in 0.5% w/v aqueous sodium carboxymethylcellulose (SCMC), was administered to rats by intragastric intubation at dosage levels of 30, 60 or 120 mg/kg/day (5 animals per sex per dose). Treatment was carried out once daily for twenty-eight consecutive days. Control animals similarly received 0.5% SCMC (10 ml/kg/day). Anatomic and clinical pathology evaluations were conducted at the end of treatment period. One male rat receiving 120 mg/kg/day was found dead on Day 20. During the third week of treatment, an ungroomed appearance and hunched posture were observed for all rats receiving 60 or 120 mg/kg/day. These clinical signs tended to persist to the study termination. During Weeks 3 and/or 4, increased salivation, of one to three days duration, was also observed for two female rats receiving 60 mg/kg/day and for all rats receiving 120 mg/kg/day. There were no changes in body weight or food consumption parameters at any dose level. In comparison with control animals, significantly higher mean corpuscular volume values were recorded for male rats receiving 120 mg/kg/day. Since the magnitude of change was minimal, no correlative changes in other hematology parameters, no changes seen in females, and individual MCV values were generally similar to those of the control values, the change in MCV values was not considered to be of toxicological importance. Significantly higher globulin levels, resulting in significantly higher total protein levels and significantly lower albumin/globulin ratios were recorded for male rats receiving 120 mg/kg/day in comparison with controls. Minor changes in male and female albumin levels and female A/G ratios were also recorded for rats in the high dosage group. In these instances, however, the apparent changes were very low in magnitude and, although statistically significant, were not considered to be of toxicological importance. Urea nitrogen levels for both male and female rats receiving 120mg/kg/day were significantly higher than those of control animals receiving 0.5% SCMC. Alkaline phosphatase and glutamic-pyruvic transaminase (GPT) levels for female rats receiving 120 mg/kg/day and GPT levels for female rats receiving 60 mg/kg/day were significantly higher than those of the corresponding control animals. The above changes in biochemical parameters were considered possibly to be related to test substance treatment. Slightly lower sodium levels were recorded for male rats receiving the test substance with statistical significance being achieved for all treatment groups. No other biochemical changes were observed. In comparison with control animals significantly higher adjusted kidney weights were recorded for both male and female rats receiving 120 mg/kg/day. Adjusted liver weights for female rats in the high dosage group were similarly significantly higher than those of the controls. For male rats receiving 60 or 120 mg/kg/day, significantly higher testis weights were recorded in comparison with

controls. The shift in this organ weight however was low in magnitude and the apparent trend to higher testes weights was not considered to be treatment-related. No histopathological changes were noted in the study.

Biodegradation test

The purpose of the test was to determine the biodegradation of test substance by closed bottle and measured dissolved oxygen. The test concentration was 2 mg/L. The preparation of the stock solution was completed by inoculating the test substance with the activated sludge at the rate of 1 drop of inoculum per liter. The concentration for the standard (Sodium Benzoate) was 3 mg/L. The test was conducted at 21±1°C and Dissolved Oxygen concentrations were determined in duplicate at 0, 5, 15 and 28 days. Oxygen depletion in 28 days was 0.475 mgO2/L. The Theoretical oxygen demand (ThOD) was 5.5 mgO2/L which resulted in a percentage biodegradation of the test substance in 28 days between 6 -13% degradation. The standard, Sodium Benzoate, had degradation of 92% in 28 days.

This information is submitted in accordance with current guidance issued by EPA indicating EPA's interpretation of Section 8(e) of the Toxic Substances Control Act or, where it is not clear that reporting criteria have been met, it is submitted as a precautionary measure and because it is information in which EPA may have an interest.

Sincerely,

S. Satheesh Anand, Ph.D., DABT
Senior Research Toxicologist

SSA/NLB: clp
(302) 366-5314

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